DATE **ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2 Exhibit)** February 2000 **BUDGET ACTIVITY** PE NUMBER AND TITLE 0602787A Medical Technology 2 - Applied Research FY 2003 FY 1999 FY 2000 FY 2001 FY 2002 FY 2004 FY 2005 Cost to **Total Cost** COST (In Thousands) Estimate Estimate Estimate Estimate Complete Actual Estimate Estimate Continuing Total Program Element (PE) Cost Continuing A838 Neurotoxin Exposure Treatment A841 Minimally Invasive Surgery A843 Health Technology Roadmaps A845 Bone Disease Research A863 Battlefield Surgical Replacement Telemedicine/Advanced Technology Continuing Continuing A870 DoD Medical Defense Against Infectious Diseases Continuina Continuina A872 Neurofibromatosis Research HIV Exploratory Research Continuing Continuing A874 Combat Casualty Care Technology Continuing Continuina A878 Health Hazards of Military Materiel Continuina Continuina Medical Factors Enhancing Soldier Effectiveness Continuing Continuing A921 Ovarian Cancer Research A948 Portable Cardiopulmonary Bypass Pump and Oxygenator **Advanced Cancer Detection** A949 A950 Teleradiology

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DATE **ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2 Exhibit)** February 2000 BUDGET ACTIVITY PE NUMBER AND TITLE 2 - Applied Research 0602787A Medical Technology FY 2001 FY 2003 FY 2004 FY 1999 FY 2000 FY 2002 FY 2005 Cost to **Total Cost** COST (In Thousands) Actual Estimate Estimate Estimate Estimate Estimate **Estimate** Complete A951 Diagnostic and Surgical Breast Imaging 1926 1926 A952 Musculoskeletal Injuries 1926 5885 0 7811 Disaster Relief and Emergency Medical Services 9809 0 0 9630 0 0 0 19439 n A962 Polynitroxylated Hemoglobin 1962 1962 A963 National Medical Testbed 14714 0 0 14714 Infomatics-based Medical Emergency Tools 4414 0 n 0 n 4414 A965 Eye Research 1962 1962 A966 Blood Research 5395 5395 Dye Targeted Laser Fusion 2943 0 0 0 0 O 0 2943 Synchrotron-based High Energy Radiation Beam 4905 0 4905

A. <u>Mission Description and Budget Item Justification</u>: The primary goal of medical research and development is to sustain medical technology superiority to improve the protection and survivability of U.S. forces on conventional battlefields as well as in potential areas of low intensity conflict and military operations short of war. This program element funds applied research in Department of Defense (DOD) medical protection against naturally occurring diseases of military importance and combat dentistry, as well as applied research for Department of Army care of combat casualties, health hazard assessment of military materiel, and medical factors enhancing soldier effectiveness. This program element is the core DOD technology base to develop methods and materials for infectious disease prevention and treatment including vaccines, prophylactic and therapeutic drugs, insect repellents, and methods of diagnosis and identification of naturally occurring infectious diseases; prevention and treatment of combat maxillofacial (face and neck) injuries, and essential dental treatment on the battlefield; combat casualty care of trauma and burns due to weapons, organ system survival, shock resulting from blood loss and infection, blood preservation, and potential blood substitutes for battlefield care; assessment of the health hazards of military materiel, and the sustainment or enhancement of soldier performance. The work in this PE is consistent with the Army Science and Technology Master Plan, Army force modernization plans, and Project Reliance. This program is managed primarily by the U.S. Army Medical Research and Materiel Command.

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A977 Emerging Infectious Diseases

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DATE **ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2 Exhibit)** February 2000 BUDGET ACTIVITY PE NUMBER AND TITLE 0602787A Medical Technology 2 - Applied Research **B. Program Change Summary** FY 1999 FY 2001 FY 2000 Previous President's Budget (FY 2000/2001 PB) 70136 138264 68014 Appropriated Value 139255 176636 Adjustments to Appropriated Value a. Congressional General Reductions -991 b. SBIR / STTR -3209 c. Omnibus or Other Above Threshold Reductions -652 d. Below Threshold Reprogramming -501 -552 Rescissions -1785 Adjustments to Budget Years Since FY 2001/2001 PB +7715 Current Budget Submit (FY 2001 PB) 134022 174199 75729

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ARMY RDT&E BUDGET ITE	ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2A Exhibit)								
BUDGET ACTIVITY 2 - Applied Research			NUMBER AND 1602787A		Technolo _s	gy			PROJECT A838
COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate	-	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost
A838 Neurotoxin Exposure Treatment	19261	98	809 0	0	0	0	0	0	29070

Mission Description and Justification: By Congressional direction, the purpose of this project is to conduct a research program on pathophysiology and treatment of neurodegenerative diseases, including Parkinson's Disease, including environmental and stress-exposure factors encountered in military operations that may be neurotoxic or lead to neurodegenerative diseases. An improved understanding of the pathophysiology of neurodegenerative diseases will form the basis of potential preventive measures against the effects of military threat agents and military operational hazards, and also lead to treatment interventions for Parkinson's Disease.

FY 1999 Accomplishments:

19261

- Completed identification of research areas most relevant to neurodegenerative risk, including acquired Parkinsonism, and initiated studies to clarify etiologies, pathologies, and therapeutic strategies (metabolic interventions, neuroprotectants, and restorative therapies) most likely to yield tangible results.

Total 19261

FY 2000 Planned Program:

- 9545 Expand and continue the program in these six thrust areas:
 - Conduct a strong basic research program to understand the fundamental nature of neural cell death and dysfunction underlying neurodegenerative diseases.
 - Identify protective agents that may be useful in neural cell dysfunction.
 - Develop improved methods for early detection of neurodegenerative disease.
 - Explore feasibility of new therapeutic strategies for neurodegenerative disease involving transplantation and neuroprotection.
 - Explore feasibility of new therapeutic strategies for neurodegenerative disease involving gene replication and other novel methods.
 - Investigate environmental factors that may be associated with neurodegenerative diseases.
- 264 Small Business Innovative Research/Small Business Technology Transfer Research Programs.

Total 9809

FY 2001 Planned Program: Project not funded in FY 2001.

Project A838 Page 4 of 40 Pages Exhibit R-2A (PE 0602787A)

ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2A Exhibit)									February 2000			
BUDGET ACTIVITY 2 - Applied Research			E NUMBER / 0602787		TITLE Medical T	echnolo	gy			PROJECT A841		
COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate			FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost		
A841 Minimally Invasive Surgery	11079	98	309	0	0	0	0	0	0	20888		

<u>Mission Description and Justification</u>: By Congressional direction, this program supports continuation of development of sophisticated computer-based surgery devices. This program will improve technologies developed under the Computer Assisted Minimally Invasive Surgery (CAMIS) program, including integration of an intraoperative ultrasound imaging device, a small fiber endoscope, and application of an intraoperative magnetic resonance imaging device.

FY 1999 Accomplishments:

• 11079 - Developed minimally invasive surgical technologies at the Center for Minimally Invasive Technology (CMIT) at Massachusetts General Hospital.

Total 11079

FY 2000 Planned Program:

- 9545 Continue development of minimally invasive surgical technologies at CMIT at Massachusetts General Hospital.
- 264 Small Business Innovative Research/Small Business Technology Transfer Research Programs.

Total 9809

FY 2001 Planned Program: Project not funded in FY 2001.

Project A841 Page 5 of 40 Pages Exhibit R-2A (PE 0602787A)

ARMY RDT&E BUDGET IT	TEM JUS	TIFIC	ATION (R	-2A Exh	ibit)		DATE Fe	bruary 20	000	
BUDGET ACTIVITY 2 - Applied Research	PE NUMBER AND TITLE 0602787A Medical Technology							PROJEC A843		
COST (In Thousands)	FY 1999 Actual	FY 200 Estimat		FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cos	
A843 Health Technology Roadmaps	1925		0 0	0	0	0	0	0	192	
FY 1999 Accomplishments: • 1925 - Completed, at the Department of Ener of advanced technology research and de - Developed a methodology for determi infrastructure costs. - Demonstrated cost reduction potential Total 1925 FY 2000 Planned Program: Project not funded in FY 2000. FY 2001 Planned Program: Project not funded in FY 2001.	evelopment pro ning medical a and information	ograms. applicatio	ons for which te	chnology car	drive down	n Department	t of Defense		_	

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Project A843

ARMY RDT&E BUDGET ITE	ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2A Exhibit)									000
BUDGET ACTIVITY 2 - Applied Research			_	MBER AND 2787A		echnolo	gy			PROJECT 4845
COST (In Thousands)	FY 1999 Actual	FY 200 Estimat	-	FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost
A845 Bone Disease Research	2408	6	6404	0	0	0	0	0	0	8812

Mission Description and Justification: This program is intended to advance bone physiology research that may lead to strategies to improve bone health of young men and women, thereby enhancing military readiness by reducing the incidence of stress fracture during physically intensive training, and reducing the incidence of osteoporosis later in life. Individual health habits that can be encouraged in young recruits may have significant effects on achievement of peak bone mineral accretion and affect other aspects of short- and long-term bone health. Understanding bone remodeling processes triggered by physical training and the relationship to injury susceptibility will reveal appropriate training and other interventions that can reduce bone injuries in military personnel. Identification of predictors of stress fracture susceptibility, efficacious interventions, and treatment strategies for susceptible and injured service members can further reduce the impact of stress fractures on readiness. The ultimate benefits of this program include establishing optimal approaches to bone health of importance to all young Americans, reduction in lost duty time from skeletal injuries, and significant medical cost avoidance for the Department of Defense and the Department of Veterans Affairs. This program fills a specific and previously neglected niche in bone physiology research, supporting a wide range of basic science through applied clinical studies on biomechanical stress on the skeleton. This is also likely to leverage related areas of importance to the military such as muscle remodeling and it supports researchers who can address other questions fundamental to bone physiology and the understanding of bone diseases; research into the pathogenesis of bone diseases substantially supports understanding of normal processes.

FY 1999 Accomplishments:

2408 - Determined initial populations at risk for increased bone injuries impacting military readiness.

- Conducted studies to clarify individual risk factors for stress fractures, restorative interventions (nutritional, training, and medical treatment) and for predictive biomarkers for field assessment of metabolic status and impending injury.
- Laid foundation for longitudinal study of military risk factors for bone injury and body composition changes.

Total 2408

FY 2000 Planned Program:

• 6231 Expand and continue the program in these six thrust areas:

- Conduct a strong basic research program to understand the fundamental nature of mechanical influences on bone cells.
- Develop methodology to overcome technological barriers in imaging that will enable sequential studies of functional changes in bone.
- Define the role of bone remodeling in stress fracture pathogenesis to determine if it would be beneficial or harmful to block remodeling in recruit training.
- Investigate interventions (e.g., calcium-nutrient drinks, weak androgens, oral contraceptives) to improve bone health in men and/or women.
- Describe changes in bone density and health in longitudinal studies of young men and women engaged in demanding training program.
- Investigate treatments that increase rates of healing after stress fracture.
- Small Business Innovative Research/Small Business Technology Transfer Research Programs.

Total 6404

Project A845 Page 7 of 40 Pages Exhibit R-2A (PE 0602787A)

ARMY RDT&E BUDGET ITEM JUS		DATE February 2000
BUDGET ACTIVITY 2 - Applied Research	PE NUMBER AND TITLE 0602787A Medical Technology	PROJECT A845
FY 2001 Planned Program: Project not funded in FY 2001.		
		- III
Project A845	Page 8 of 40 Pages	Exhibit R-2A (PE 0602787A)

ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2A Exhibit)									000
BUDGET ACTIVITY 2 - Applied Research			NUMBER AND 602787A		echnolo	gy			PROJECT A863
COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate		FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost
A863 Battlefield Surgical Replacement	0	24	52 0	0	0	0	0	0	2452

<u>Mission Description and Justification</u>: By Congressional direction, the purpose of this program is to establish a process to select medical research projects of clear scientific merit and direct relevance to military health including tissue regeneration for combat casualty care.

FY 1999 Accomplishments: Project not funded in FY 1999.

FY 2000 Planned Program:

- 2386 Awaiting proposal submission for evaluation to be followed by contract award.
- Small Business Innovative Research/Small Business Technology Transfer Research Programs.

Total 2452

FY 2001 Planned Program: Project not funded in FY 2001.

Project A863 Page 9 of 40 Pages Exhibit R-2A (PE 0602787A)

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		ARMY RDT&E BUDGET	ITEM JUS	TIFICAT	TION (R-	-2A Exh	ibit)		DATE Fe	bruary 20	000
BUDGET ACT 2 - Appli		search			UMBER AND 02787A I	TITLE Viedical T	PROJECT A869				
		COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate	FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cos
A869 Telen	medicine/A	dvanced Technology	3183	5213	4467	4480	3306	3499	3566	Continuing	Continuir
application of system to de capable of fi	of physio etermine functionin nications in ccomplis	 Began to modify the Land Warrior Evaluated a miniaturized eye oxime microimpulse radar unit to assess care Supported Joint Medical Operations 	(biophysical and nelude developing on identification, and telerobotics) System to allow verter to assess cere diovascular functs. Telemedicine A rfighter Physiology, core and sking ogy development	biochemical ig the ability and initial of s. wound detect bral blood of ion. dvanced Co gical Status temperature of a flexible	I sensors and to quickly a development ation and rem exygen contection Monitoring geolocation e ureteroscop	fusion). Reund accurated of parallel a note triage count for measurology Demo (WPSM) syst) that collect simulator	search will f y determine and supporting mmunication res of brain p instration. tem for use a s and stores with various	ocus on devouchen a sold g technologien between in berfusion. E	eloping a we ier is minima ies and system adividual solo explored use ounted Battles in an open, s	arable, integ ally impaired ms, including diers and the of a miniatur space Battle standardized	medic. ized Lab that format.
Total	3183										
FY 2000 P	Planned I 928	- Investigate an advanced pulse oxim	etry signal analys	sis capability	y as a compo	ment of a rea	l-time soldie	r monitoring	g system. Be	-i1ti	
		noninvasive physiologic monitoring									
•	1166	noninvasive physiologic monitoring scardiac output monitoring. - Evaluate acoustic methods to diagnoblood oxygen saturation and organ peclosed head trauma.	system for use by ose tension pneur	medics in the	he battlefield on the battlef	l. Continue of the field. Test a	exploration of first-generat	of a microim ion eye oxin	pulse radar uneter to non-	init for nonir invasively m	easure
•	1166 2239	cardiac output monitoring. - Evaluate acoustic methods to diagnoblood oxygen saturation and organ per closed head trauma. - Test first-generation WPSM for phy collect mission-specific physiologica	system for use by ose tension pneur erfusion. Begin it rsiological monited data from soldie	medics in the mothoraces of needing of solders during fields.	he battlefield on the battlef into a nonin lier status. I eld testing.	I. Continue of the continue of	exploration of first-generateranial pressu SM system v	of a microim ion eye oxin are monitor	pulse radar u neter to non- to assess intr	init for nonir invasively m acranial pres	easure sure in
•		cardiac output monitoring. - Evaluate acoustic methods to diagnoblood oxygen saturation and organ peclosed head trauma. - Test first-generation WPSM for phycollect mission-specific physiologica - Support Joint Medical Operations-T	system for use by ose tension pneur erfusion. Begin it rsiological monited data from soldical delemedicine Adv	medics in the mothoraces of the nothoraces of the nother state of	he battlefield on the battlef into a nonin lier status. I eld testing. cept Technology	I. Continue of the continue of	exploration of first-generateranial pressu SM system value tration.	of a microim ion eye oxin are monitor	pulse radar u neter to non- to assess intr	init for nonir invasively m acranial pres	easure sure in
•	2239 740 140	cardiac output monitoring. - Evaluate acoustic methods to diagnoblood oxygen saturation and organ per closed head trauma. - Test first-generation WPSM for phy collect mission-specific physiologica	system for use by ose tension pneur erfusion. Begin it rsiological monited data from soldical delemedicine Adv	medics in the mothoraces of the nothoraces of the nother state of	he battlefield on the battlef into a nonin lier status. I eld testing. cept Technology	I. Continue of the continue of	exploration of first-generateranial pressu SM system value tration.	of a microim ion eye oxin are monitor	pulse radar u neter to non- to assess intr	init for nonir invasively m acranial pres	easure sure in
• • • Total	2239 740	cardiac output monitoring. - Evaluate acoustic methods to diagnoblood oxygen saturation and organ peclosed head trauma. - Test first-generation WPSM for phycollect mission-specific physiologica - Support Joint Medical Operations-T	system for use by ose tension pneur erfusion. Begin it rsiological monited data from soldical delemedicine Adv	medics in the mothoraces of the nothoraces of the nother state of	he battlefield on the battlef into a nonin lier status. I eld testing. cept Technology	I. Continue of the continue of	exploration of first-generateranial pressu SM system value tration.	of a microim ion eye oxin are monitor	pulse radar u neter to non- to assess intr	init for nonir invasively m acranial pres	easure sure in

	-	ARMY RDT&E BUDGET	ITEM JUSTIFICATION (R-2A Exhibit)	DATE February 2000
BUDGET AC 2 - App	CTIVITY lied Res	search	PE NUMBER AND TITLE 0602787A Medical Techi	PROJECT nology A869
Y 2001	Planned P	rogram:		
•		- Continue testing noninvasive sense	ors for Warrior Medic to assist in far-forward remote triage. gent instructional systems to facilitate adaptive learning.	Continue evaluating Warrior Medic and WPSM
•	927		acquisition and management capabilities, to support the form	ulation and testing of modeling strategies to predict
•	1504		cal technology overlays to tactical computing/communicating ary to pre-injury. Test artificial intelligence/sensor fusion pro	
Total	4467	injury and to compare data post-inju	iry to pre-injury. Test artificial intelligence/sensor fusion pro	tocols for WPSM.
Project A8	869		Page 11 of 40 Pages	Exhibit R-2A (PE 0602787A)

ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2A Exhibit)									000
BUDGET ACTIVITY 2 - Applied Research		E NUMBER AN 0602787A	Medical	PROJECT A870					
COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate		FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost
A870 DoD Medical Defense Against Infectious Diseases	23055	236	674 248	25611	28574	30324	32178	Continuing	Continuing

<u>Mission Description and Justification</u>: This project supports development of medical countermeasures to naturally occurring infectious diseases, a significant threat to forces deployed outside the United States. These countermeasures will protect the force from infection and sustain operations by preventing hospitalizations and evacuations from the theater of operations.

FY 1999 Accomplishments:

- 2934 Conducted applied research on vaccines to prevent hepatitis E and to prevent meningitis caused by Group B meningococcus, on a diagnostic device for scrub typhus, on a natural insecticide, and on control of insect vectors of disease.
 - Conducted epidemiological studies of hepatitis E at multiple sites around the world to help assess risk to deployed military forces.
 - Conducted the first human evaluation of a vaccine to prevent hepatitis E.
 - Evaluated a rapid test to detect scrub typhus infection.
 - Conducted animal studies to compare two vaccine candidates for prevention of Group B meningococcal infection needed for further down-selection prior to human clinical studies.
 - Characterized a candidate insecticide that incorporates a chemical naturally produced by bacteria that is toxic to insects.
 - Characterized insect populations and bite rates at field sites for vaccine and drug studies.
- 2201 Conducted applied research on vaccines to prevent the most common causes of bacterial diarrhea.
 - Modified Shigella vaccine candidate antigens to try to improve their safety.
 - Evaluated the efficacy of combined *Shigella flexneri* 2a and *Shigella sonnei* vaccine in an animal model and demonstrated feasibility of protection from diverse *Shigella* types using a combination vaccine.
 - Studied proteins involved in the ability of Campylobacter to produce diarrheal disease, providing a systematic rationale for vaccine development.
 - Conducted studies to define which toxins and other bacterial factors are associated with disease caused by enterotoxigenic *Escherichia coli* (ETEC), important for design of vaccine candidates.
 - Established laboratory assays to measure intestinal immune response to candidate ETEC vaccines.
 - Identified four new colonization factors that may be important for ultimate development of a protective ETEC vaccine.
 - Cloned and expressed a colonization factor, CS3, for use in an ETEC vaccine based on multiple colonization factors.
 - Completed preclinical studies of CS6 (a candidate ETEC vaccine) necessary for an Investigational New Drug (IND) application to the Food and Drug Administration and ultimate clinical testing.
- 2840 Conducted applied research on vaccines to prevent viral diseases capable of interrupting combat operations.
 - Evaluated the effect of adding compounds to candidate dengue vaccines that target immune cells, necessary for designing optimal vaccine systems for stimulation of a protective immune response in recipients.

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ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2A Exhibit) BUDGET ACTIVITY 2 - Applied Research PE NUMBER AND TITLE 0602787A Medical Technology PE NUMBER AND TITLE 0602787A Medical Technology PE NUMBER AND TITLE 0602787A Medical Technology

FY 1999 Accomplishments: (continued)

- Made and characterized two candidate DNA vaccines for two of the four serotypes of dengue.
- Demonstrated the efficacy of candidate vaccines against hantaviruses in an animal model.
- In animal models, demonstrated the safety and the capability to produce an immune response of candidate vaccines against Lassa Fever, Crimean-Congo Hemorrhagic Fever and tickborne encephalitis viruses.
- 4025 Conducted applied research on candidate vaccines for prevention of malaria and on antimalarial drugs to prevent or treat malaria.
 - Conducted immunological studies to identify components of the malaria parasite to be used in engineered vaccines, either protein or DNA.
 - Used a common bacteria to produce three malaria proteins to use in evaluating the immune response to naturally occurring malaria infections or to use in vaccines to protect against malaria.
 - Studied novel routes of immunization with malaria protein and DNA vaccines to determine the impact of immunization route on immune response in the recipient.
 - Developed improved methods for immunization with DNA-based vaccines, important for eliciting a more effective immune response in recipients.
 - Established and validated a rhesus monkey model for malaria for assessing vaccine strategies for malaria.
 - Demonstrated that synthetic compounds of nucleic acids (oligodeoxynucleotides or ODNs) of specific composition are potent inducers of nonspecific immunity and excellent additives for enhancing the immunizing effects of protein-based vaccines.
 - Expressed and purified recombinant proteins for five different target proteins for structure-based drug design of novel antimalarial drugs, a necessary step in the rational design of new candidate drugs.
 - Expanded existing capabilities to screen antimalarial drugs by developing new animal models. Analyzed the antimalarial activity of novel candidate compounds.
 - Developed tests to monitor the development and spread of drug-resistant malaria, important for guiding new drug development and the use of currently available drugs.
- 11055 Paid administrative overhead costs at the Walter Reed Army Institute of Research (WRAIR).

Total 23055

FY 2000 Planned Program:

- 4051 Conduct applied research on candidate vaccines for prevention of malaria and on antimalarial drugs to prevent or treat malaria.
 - Develop standardized methods to reliably measure immune responses to candidate malaria vaccines.
 - Conduct preclinical studies of candidate vaccines to support an IND application.
 - Develop a method to perform human experiments where the *Plasmodium vivax* parasite can be introduced into human volunteers to test the ability of candidate vaccines to prevent disease caused by *Plasmodium vivax*, the second-most important cause of malaria.
 - Synthesize candidate antimalarial drugs or isolate candidate drugs from natural products.
 - Develop techniques for the cultivation and drug sensitivity testing of vivax malaria.
 - Express target proteins for structure-based drug design and determine modes of action and resistance of parasites to antimalarial drugs.
 - Create a deployable field test to assay drug sensitivity patterns in malaria for use in monitoring parasite development of drug resistance.

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DATE **ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2A Exhibit)** February 2000 PE NUMBER AND TITLE BUDGET ACTIVITY **PROJECT** 2 - Applied Research 0602787A Medical Technology A870 FY 2000 Planned Program: (continued) - Conduct screening to measure activity or cytotoxicity of candidate drugs. - Prepare radiolabelled drug candidates for preclinical studies of drug distribution, pharmacokinetics, and metabolism. - Perform preclinical toxicology studies of new drugs. 2254 Conduct applied research on vaccines to prevent the most common causes of bacterial diarrhea. - Modify candidate live Shigella vaccines to reduce vaccine-induced toxicity and/or fecal excretion vaccine organisms while retaining efficacy in prevention of diarrhea. - Improve candidate live vaccines so that orally administered vaccine organisms can be rapidly identified if they are excreted. - Devise polyvalent vaccines so that service members can be protected against the many different types of Shigella bacteria that could cause diarrhea. - Characterize ETEC virulence factors to find new potential vaccine components. - Devise methods to boost mucosal immune responses to oral vaccines. - Develop an improved animal model for ETEC infection to enable testing of vaccine candidates and prediction of efficacy in animals. - Improve methods to diagnose ETEC infections, which are needed for testing efficacy of the candidate vaccines in humans. - Explore new and/or improved animal models of *Campylobacter* enteritis and immunity, including the ferret, the pig, and nonhuman primates. - Improve methods to diagnose *Campylobacter* infections, which are needed for testing efficacy of candidate vaccines in humans. 3045 Conduct applied research on the components of diagnostic tests to be applied to a common diagnostic device for biological defense and infectious disease threats; on vaccines to prevent hepatitis E, scrub typhus, and meningitis caused by Group B meningococcus; and on control of insect vectors of disease. - Develop infectious disease-specific reagents for malaria, enteric diseases, dengue viruses, and the hemorrhagic fever viruses so they are compatible for use on the portable Common Diagnostic Platform for Biological Defense and Endemic Infectious Diseases capable of detecting and identifying nucleic acids. - Establish the magnitude of the immune response to hepatitis E virus (HEV) antibody that is necessary to prevent disease. - Further characterize the human immune responses to HEV infection, disease, and vaccine. - Refine epidemiology of HEV in Asia and Africa. - Sustain or refute presence of hepatitis E disease among humans in Latin America. - Characterize the animal host (particularly rodents) of HEV and the HEV isolates obtained. - Determine the feasibility of vaccine development against scrub typhus. - Optimize Group B meningitis candidate vaccine strains. - Complete animal immunogenicity and safety studies to determine the optimal presentation and formulation of Group B meningitis vaccine. - Conduct a detailed analysis of the animal and human immune responses to the Group B meningitis candidate vaccines to determine the best antigens. - Establish a standard insecticide resistance and susceptibility test and test insects capable of transmitting disease to determine if the military insecticides are still effective. - Evaluate the threat of tick- and chigger-borne diseases to the U.S. military.

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FY 2000 Planned Program: (continued)

Project A870

	-	ARMY RDT&E BUDGET IT	EM JUSTIFICATION (R-2A Exhib	oit) Pebruary 2000
	ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2A Exhibit) PENUMBER AND TITLE 0602787A Medical Technology - Begin development and evaluation of a dengue mosquito vector control system consisting of integrated packaged for a preventive medicine detachment (or service equivalent). - Conduct preliminary development of devices and techniques that may serve as components of a vector of malaria, including a field device for detecting Plasmodium in mosquitoes. 2844 Conduct applied research on vaccines to prevent viral diseases capable of interrupting combat operations. - Test candidate dengue vaccines to determine if they will be effective in protecting recipients of diverse good and to be a validate a method for quantifying antibodies that exacerbate dengue disease. - Characterize determinants leading to severe dengue disease. - Characterize mechanisms of viral hemorrhagic fever (VHF) and encephalitis pathogenesis. - Develop candidate VHF and encephalitis vaccines and test in animal models. - Evaluate antiviral drug candidates for efficacy in vitro and in animal models. - Develop and evaluate primate monoclonal antibodies for passive protective efficacy in animal models in Improve capability to rapidly identify VHF and encephalitis agents in the field and to provide definitive - Develop candidate vaccines against one or more hantaviruses and test in animals to assess immune respective expensive provential and animals and also isonal numan monoclonal antibodies or antiviral drugs) and testing them in cell culture and animals and also isonal numan monoclonal antibodies or antiviral drugs) and testing them in cell culture and animals and also isonal part of the programs.		PROJECT	
•		- Begin development and evaluation of packaged for a preventive medicine deta - Conduct preliminary development of community malaria, including a field device for deta - Conduct applied research on vaccines to - Test candidate dengue vaccines to deta - Validate measures of immune T cell malaria, including a field device for deta - Validate anethod for quantifying anti - Characterize determinants leading to so - Determine the feasibility of second-gearcharacterize mechanisms of viral hemalicated properties and evaluate antiviral drug candidates for - Develop and evaluate primate monoclarimprove capability to rapidly identify - Develop candidate vaccines against or - Improve capability to rapidly identify,	a dengue mosquito vector control system consisting of achment (or service equivalent). devices and techniques that may serve as components ecting <i>Plasmodium</i> in mosquitoes. The prevent viral diseases capable of interrupting combast ermine if they will be effective in protecting recipient memory and assess the relevance of these immune cell abodies that exacerbate dengue disease. The energy disease in the field should be dengue vaccines. The energy in vitro and in animal models, and antibodies for passive protective efficacy in anim VHF and encephalitis agents in the field and to provide or more hantaviruses and test in animals to assess in assess risk, and formulate control strategies for hantales.	of integrated tools and information that can be physically sof a vector control system for mosquitoes that transmit at operations. Its of diverse genetic backgrounds. Its to protection against dengue disease. In all models including primates. It de definitive confirmation in reference labs. It immune responses and protection. It is a protection against dengue disease.
•	11056			and also isolating and characterizing novel nantaviruses.
• Total		- Small Business Innovative Research/S	small Business Technology Transfer Research Program	ms.
FV 2001		rogram.		
•		Conduct applied research on candidate - Conduct preclinical studies of DNA va - Express proteins encoded by the <i>Plast</i> an immune response in an animal mode - Develop field sites for <i>P. vivax</i> human - Develop a field site for testing a drug - Determine a strategy to render the <i>P. f</i> Conduct applied research on vaccines to	accines to prevent <i>P. falciparum</i> malaria. modium vivax gene homologs of the <i>P. falciparum</i> car el. a vaccine trials. for treatment of multidrug-resistant malaria. falciparum multidrug-resistant gene ineffective. b prevent the most common causes of bacterial diarrho	ndidate vaccine components and test their ability to induce
l 		- Complete animal trials of candidate <i>S</i> .	dysenteriae vaccines.	
FY 2001	I Planned I	Program: (continued) - Characterize enteric bacterial proteins	identified through genomic sequence data analysis to	assess their possible application to vaccine development.
Project A	A870		Page 15 of 40 Pages	Exhibit R-2A (PE 0602787A)

	ARMY RDT	&E COST ANALYSIS (R-3)	DATE February 2000
BUDGET ACTIVITY 2 - Applied R		PE NUMBER AND TITLE 0602787A Medical Technol	PROJECT A870
306273	 Characterize the optimal formulating and the continuous of the evaluation of the evaluation of the conduct applied research on the condisease threats; on vaccines to prevent disease. Transition components of the DNA disease threats. Assess the threat of hepatitis E to be disease threats. Demonstrate the feasibility of immit vaccine in an animal model. Genetically alter the Group B menerolevelop a rapid immunological menerolevelop a rapid immunological menerolevelop and the conduct applied research on vaccines. Assess the threat of hemorrhagic ferolevelop a cytotoxic T cell technolevelop and cytotoxic T cell technolevelo	es associated with recovery from <i>Campylobacter</i> infection and semponents of diagnostic tests to be applied to a common diagnostent hepatitis E, scrub typhus, and meningitis caused by Group E. A tests for malaria and dengue to functional use on the common U.S. service members in Africa and Latin America. Interpolation against scrub typhus in an animal model and interpolation against scrub typhus in an animal model ani	stic device for biological defense and infectious B meningococcus; and on control of insect vectors on diagnostic platform for biodefense and infectious and demonstrate efficacy of a candidate scrub typhus roduce it in a vaccine manufacturing process. Trations. Truses.
• 1105 Total 2484	.,	at WRAIR.	
Project A870		Page 16 of 40 Pages	Exhibit R-3 (PE 0602787A)

ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2A Exhibit)									February 2000			
		_			echnolo	gy			PROJECT A872			
FY 1999 Actual			-	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost			
11079	147	714	0	0	0	0	0	0	25793			
	FY 1999 Actual	FY 1999 FY 2000 Actual Estimate	PE NUMBER A 0602787 FY 1999 FY 2000 FY 200 Estimate Estimate	PE NUMBER AND 0602787A N FY 1999 FY 2000 FY 2001 Estimate Estimate	PE NUMBER AND TITLE 0602787A Medical T FY 1999 FY 2000 FY 2001 FY 2002 Actual Estimate Estimate Estimate	PE NUMBER AND TITLE 0602787A Medical Technology FY 1999 FY 2000 FY 2001 FY 2002 FY 2003 Actual Estimate Estimate Estimate Estimate	PE NUMBER AND TITLE 0602787A Medical Technology FY 1999 FY 2000 FY 2001 FY 2002 FY 2003 FY 2004 Actual Estimate Estimate Estimate Estimate Estimate	PE NUMBER AND TITLE 0602787A Medical Technology FY 1999 FY 2000 FY 2001 FY 2002 FY 2003 FY 2004 FY 2005 Actual Estimate Estimate Estimate Estimate Estimate Estimate	PE NUMBER AND TITLE 0602787A Medical Technology FY 1999 FY 2000 FY 2001 FY 2001 FY 2002 FY 2003 FY 2004 FY 2005 Estimate FY 2005 Estimate FY 2006 FY 2007 Estimate FY 2007 FY 2008 FY 2008 FY 2009 F			

<u>Mission Description and Justification</u>: By Congressional direction, the purpose of this appropriation is only for neurofibromatosis research.

FY 1999 Accomplishments:

11079 - Received 21 proposals in October 1998 for the FY 1998 program.

- Completed peer and programmatic review by April 1999. Awarded nine grants. Received 2-year funds in January 1999 for the FY 1999 program.
- Held vision setting meeting in April 1999. Published a program announcement in June 1999 and received 48 proposals in September 1999.

Total 11079

FY 2000 Planned Program:

- Complete peer and programmatic review by January 2000 and negotiate awards by September 2000 for the FY 1999 program. Receive funds in January 2000, hold vision setting meeting in January 2000, and publish a program announcement in February 2000 for the FY 2000 program. Receive proposals in May 2000 and conduct peer review in August 2000.

396 - Small Business Innovative Research/Small Business Technology Transfer Research Programs.

Total 14714

FY 2001 Planned Program: Project not funded in FY 2001. Programmatically review and award proposals from the FY 2000 program.

Project A872 Page 17 of 40 Pages Exhibit R-2A (PE 0602787A)

ARMY RDT&E BUDGET ITE	ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2A Exhibit)								
BUDGET ACTIVITY 2 - Applied Research PE NUMBER AND TITLE 0602787A Medical Technology							PROJECT A873		
COST (In Thousands)	(s) FY 1999 FY 2 Actual Estin			FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost
A873 HIV Exploratory Research	125	541 11579	11021	10890	11372	11586	Continuing	Continuing	

Mission Description and Justification: This project provides for applied research of improved diagnostics, epidemiology, candidate immunogens, promising drugs and behavioral modification for prevention and treatment of human immunodeficiency virus (HIV). Main efforts include developing experimental models of disease, preparation of new vaccine candidates, improved diagnosis of disease, and risk assessment. Current policy prohibits antibody-positive service members from deployment outside the continental United States. A safe and effective vaccine for prevention of infection and intervention techniques will permit all service members to become worldwide deployable.

FY 1999 Accomplishments:

- 12033 Conducted applied research on novel candidate vaccines to prevent HIV infection, on the human immune response factors that predict protection from HIV infection or disease, on human cohorts for potential testing of HIV vaccines, and on animal models for testing candidate vaccines.
 - Conducted preclinical studies of clade B oligomeric protein vaccine candidates, necessary for advancement to clinical testing.
 - Established domestic and Thai laboratory infrastructure for support of vaccine efficacy trials, important for developing and standardizing measures of vaccine performance in future clinical studies of vaccine candidates.
 - Successfully established a rhesus macaque challenge model for human HIV infection using a simian-human immunodeficiency virus chimeric virus; the model is useful for assessing immunogenicity and protective efficacy of candidate HIV envelope-based vaccines.
 - Constructed and tested a DNA vaccine and demonstrated induction of antibody that is both quantitatively and qualitatively superior to recombinant protein vaccines used alone. The findings are promising and may lead to future vaccines that induce protective responses in recipients.
 - Conducted a vaccine study in baboons that suggests cross-protection of subtype B HIV infection using a single subtype E rgp120 vaccine, important for design of vaccines capable of protecting against multiple subtypes of HIV.
 - Studied viral correlates of HIV transmission and pathogenesis, important for design of vaccines and for selection of measures to be used in clinical studies to characterize and assess vaccine effectiveness.
 - Characterized HIV-specific protective epitopes of vaccine products for national and international use, important for selection and production of candidate vaccine components.
- 1780 Paid administrative overhead costs at the Walter Reed Army Institute of Research (WRAIR).

Total 13813

FY 2000 Planned Program:

- 10432 Conduct applied research on novel candidate vaccines to prevent HIV infection, on the human immune response factors that predict protection from HIV infection or disease, on human cohorts for potential testing of HIV vaccines, and on animal models for testing candidate vaccines.
 - Evaluate the importance of HIV genotypes in predicting HIV immunotypes necessary for inclusion in an HIV vaccine.

Project A873 Page 18 of 40 Pages Exhibit R-2A (PE 0602787A)

PE NUMBER AND TITLE 2 - Applied Research Per 2000 Planned Program: (continued) - Define the correlates of immunity to HIV, necessary for vaccine design Establish genetic and phenotypic correlates of drug resistance among HIV-1 clinical isolates, necessary for establishing drug treatme military dependents Conduct animal studies of candidate HIV vaccines to prevent HIV infection Evaluate and validate a rapid test for field diagnosis of HIV infection Evaluate and validate are apid test for field diagnosis of HIV infection Paid administrative overhead costs at WRAIR 329 - Small Business Innovative Research/Small Business Technology Transfer Research Programs. FY 2001 Planned Program: - 9859 - Conduct applied research on novel candidate vaccines to prevent HIV infection, on the human immune response factors that predict predict prediction or disease, on human cohorts for potential testing of HIV vaccines, and on animal models for testing candidate vaccines - Clinically validate novel diagnostic and prognostic measurements of HIV-1 virological markers, necessary for establishing and stand measures of vaccine effectiveness for clinical efficacy studies Conduct preclinical studies of novel vaccine candidates Pay administrative overhead costs at WRAIR.	ebruary 2000	2A Exhibit)	M JUSTIFICATION (R-2	ARMY RDT&E BUDGET ITEM	Α	
 Define the correlates of immunity to HIV, necessary for vaccine design. Establish genetic and phenotypic correlates of drug resistance among HIV-1 clinical isolates, necessary for establishing drug treatme military dependents. Conduct animal studies of candidate HIV vaccines to prevent HIV infection. Evaluate and validate a rapid test for field diagnosis of HIV infection. Paid administrative overhead costs at WRAIR. Small Business Innovative Research/Small Business Technology Transfer Research Programs. PS 2001 Planned Program: Ocnduct applied research on novel candidate vaccines to prevent HIV infection, on the human immune response factors that predict prediction or disease, on human cohorts for potential testing of HIV vaccines, and on animal models for testing candidate vaccines according to the companies of vaccine effectiveness for clinical efficacy studies. Conduct preclinical studies of novel vaccine candidates. Pay administrative overhead costs at WRAIR. 	PROJECT A873			search		
Total 12541 FY 2001 Planned Program: 9859 Conduct applied research on novel candidate vaccines to prevent HIV infection, on the human immune response factors that predict program: HIV infection or disease, on human cohorts for potential testing of HIV vaccines, and on animal models for testing candidate vaccines. Clinically validate novel diagnostic and prognostic measurements of HIV-1 virological markers, necessary for establishing and standard measures of vaccine effectiveness for clinical efficacy studies. Conduct preclinical studies of novel vaccine candidates. 1720 - Pay administrative overhead costs at WRAIR.	g treatment strategies for		tes of drug resistance among HIV-1 cl vaccines to prevent HIV infection. d diagnosis of HIV infection. RAIR.	 Define the correlates of immunity to HIV, n Establish genetic and phenotypic correlates military dependents. Conduct animal studies of candidate HIV va Evaluate and validate a rapid test for field di Paid administrative overhead costs at WRAD 	1780	FY 2000 F
 9859 Conduct applied research on novel candidate vaccines to prevent HIV infection, on the human immune response factors that predict predict prediction or disease, on human cohorts for potential testing of HIV vaccines, and on animal models for testing candidate vaccines - Clinically validate novel diagnostic and prognostic measurements of HIV-1 virological markers, necessary for establishing and standard measures of vaccine effectiveness for clinical efficacy studies.		ž	27			Total
	vaccines.	es, and on animal models for testing can	ts for potential testing of HIV vaccine prognostic measurements of HIV-1 virtual efficacy studies.	HIV infection or disease, on human cohorts for Clinically validate novel diagnostic and progressives of vaccine effectiveness for clinical - Conduct preclinical studies of novel vaccine		•
			AIR.	- Pay administrative overhead costs at WRAI		• Total

	ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2A Exhibit) PE NUMBER AND TITLE PE NUMBER AND TITLE											
BUDGET AC 2 - App	CTIVITY lied Res	search			UMBER AND 02787A		PROJECT A874					
		COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate	FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost	
A874 Con	nbat Casualt	y Care Technology	10440	8537	8806	9063	10633	11456	12011	Continuing	Continuin	
soldiers wo weapons-in	ounded in conduced trav Accomplish 1000 2250	and Justification: This project funds to ombat and to support low-intensity comma and burns, and shock due to blood lowers: - Awarded contract to study pain man - Formulated a storage solution that we to Milestone 0. Tested foam-based he and functionality of platelets after liqueleukocytes and survival in hemorrhage - Established reverse transcriptase/pol Established human bronchial/tracheal monoxide poisoning. Investigated the apoptotic response of skin epithelial of the Combat Support Trauma and Transmethods to treat tension pneumothora - Supplemented Life Support for Trau - Completed study demonstrating anti Food and Drug Administration (FDA) Joint Medical Operations-Telemedicin Surgical Research.	bat as well as ross. It also fundagement to Gurill support refresonation and reinfusion ymerase chain and small airway effects of hemells to burns an sport (CSTAT ces. Investigat ma and Transpericrobial efficies approval and designation of the second statement of the se	thrie Research igerated stores in preclinic 5 days. Assent. reaction assert yepithelial ne proteins of didentified identified microencort (LSTAT) acy of coated concluded st	ch Institute for age of red blocal animal messed the effect ay for the qual cell model an intric oxide apoptotic master apoptotic master apoptotic master apoptotic master apoptotic master and platform. In the stainless study on 5% a	or research is cood cells for odels. Deve ects of hemo antification and determine levels and arkers and ki far-forward stioxidants ar eel external queous sulfar	nto sodium control 10 weeks. If and blood points and the lyzed red bloof tissue cytoned that the true cell viability netics. Control their effective carried their effective carried their effective control intensive carried their effective control to their effective control to the control to	channel protested animal cood cells on in polymorphinued develope and diagnosts on wounder an animal student in top	estigation of estimate as potent a synthesis af noke is unrelable opment of a mostic support I healing in a model of borical treatment	the treatment it is a pain moor in the pain moor in the production by the production by the production by the production in the production	dulators. ic dressing e life span y agic shock. on tudied the rsion of valuated s. Received Supported	
FY 2000 I	Planned Pr 1168	rogram: - Evaluate the enhancement of clot exp for transfusion-related multiple organ f drugs to decrease blood loss after sever Continue to develop liquid red blood co assessment of efficacy and safety of dr	ailure after tran e liver injury. ell storage syste	nsfusion of e Assess the i em to achiev	extended stor mportance o	age-life red f hypotherm	blood cells. ia as an indu	Evaluate the cer of coagu	e potential us lopathy duri	e of FDA-ap ng hemorrha	proved ge.	
Project A8	874			Page 20 oj	f 40 Pages			Exhib	it R-2A (PE	0602787A)	ı	

		ARMY RDT&E BUDGET ITEI	M JUSTIFICATION (R-2A Exhibit)	February 2000
BUDGET AC 2 - App		search	PE NUMBER AND TITLE 0602787A Medical Technol	ology A874
FY2000 I	Planned P	Program: (continued)		
•	1223	- Evaluate hypotensive resuscitation after herebleeding occurs during resuscitation in an resuscitation after combined brain trauma a Investigate the use of cytofluorometric mea	emorrhage as an optimal resuscitation strategy on the bat a aortotomy model to identify an important resuscitation and hemorrhage. Investigate methods to protect endother sures to select combined therapies to inhibit inflammatication and plasma volume following severe trauma and described the sures are selected.	parameter. Evaluate hypertonic fluid therapy for lial cell integrity after ischemia/reperfusion injury. on after hemorrhage and resuscitation. Test
•	1355	- Examine antioxidative neuroprotective eff Test n-acetylaspartylglutamate and n-acetyl (ADP-ribose) polymerase as a target for neu- lisofylline in burn patients with inhalation in protection from smoke inhalation injury in	ficacy of polynitroxyl hetastarch and the oxygen-carryin ated linked acidic dipeptidase inhibitors to protect again aroprotective therapies following traumatic brain injury. Investigate defective immune responses following human bronchial/tracheal and small airway epithelial ce NA microarray assay. Develop inhibitors of constitutive	ng red blood cell substitute polynitroxyl hemoglobin nst ischemia/reperfusion injury. Investigate poly . Undertake a pharmacodynamic study of ng exposure to heat. Establish models to examine ells. Identify and quantitate inflammatory mediator
•	4638	- Develop methods for sterilization of denta	all equipment at far-forward locations. Continue development and administrative expenses for the Institute of Soft Technology Demonstration.	
•	153		Il Business Technology Transfer Research Programs.	
Total	8537			
FY 2001 F	Planned P	rogram:		
•	1900	noninvasive physiologic monitoring system	e miniSTAT – as a far-forward intensive care and diagn for use by medics on the battlefield. Continue develop cord, and other organs. Complete preclinical evaluation	ment of in vivo models and testing of therapies for
•	1965	wound and injury repair techniques to corre	e and thermal inhalation injuries. Continue research into ect battle or training injuries. Investigate the diagnosis a simplify treatment of trauma in austere environments.	
•	1841	- Continue evaluation of techniques for the	formulation and assessment of efficacy and safety of dratic dressings. Continue testing of fibrin foam-based her	
•	3100	- Pay general and administrative expenses f		
Total	8806			
Project A8	874		Page 21 of 40 Pages	Exhibit R-2A (PE 0602787A)

	ļ	ARMY RDT&E BUDGET	TITEM JUS	TIFICAT	TION (R-	2A Exh	ibit)		DATE Fe	bruary 20	y 2000	
BUDGET AC		search			UMBER AND 02787A		PROJECT A878					
		COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate	FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost	
A878 Hea	alth Hazards	of Military Materiel	8329	9267	10642	11369	11718	12182	12700	Continuing	Continuin	
Emphasis i and training generated be non-ionizing extent of ex	is on identicle. Specification of the properties	and Justification: This project focution of health hazards inherent to chazards include repeated impact/jold eapons systems; toxic chemical hazard directed energy sources (laser); and potential hazards; delineating exposut to support protection criteria; and de hments: - Identified a safe range of helmet we performance. Discovered that helm different based on performance indicates overhydration problems. - Refined predictive model of toxic biomarker studies. - Developed and beta tested an initial and incidents. - Completed first phase of a low-con 75% (2-4 hours).	the engineering det and vibration streeds associated with a environmental street thresholds for illuveloping and valid eveights and centers are mass design criticators. Army Fluid Replacembustion gas in all version of the Landau vibration and version of the Landau vibration street mass design criticators.	esign and op- ess from the of deployment essors (e.g., llness or inju- lating model of of mass that teria for fem- acement Gui- capacitation aser Accider	erational use operation of t into environ heat, cold, to try; identifyits for hazard t can be tole; ale and male delines for h with incorporat and Incide	of equipment combat vehicements contained altiming exposure assessment, arated by femicial exposure assessment arated by femicial exposure and the contained arated by femicial exposure and the contained arated by femicial exposure are arrested as a femicial exposure are a femicial exposure	nt, systems a cles and airc aminated wit tude). Speci thresholds for injury predict ale helicopted viators are the raining to en-	and materiel craft; blast over hindustrial fic research for performanction, and he er pilots with the same base sure preventation fire supplicate query symmetrical preventation for the supplication of the s	used in Army verpressure a waste and ag tasks include nee degradati alth and perf nout affecting ed on head mand ion of dehyde pressant alter	y combat op nd impulse i ricultural ch e characteriz on; establish formance pro- g health or notion but m ration witho natives and	erations noise noise nemicals; ing the ning otection. ay be ut causing injury or accidents	
FY 2000 F	Planned Pr 2498	rogram: - Study injury mechanisms for exportance operational exposure limits.	osure conditions inl	herent to mil	litary lasers o	operating in	he visible ar	nd near infra	red region of	the spectru	m to refine	
•	1473 3242		ucts, and select mix gy of combined fir	xtures, for the	eir ability to ure into com	cause birth	defects.		•			

	-	ARMY RDT&E BUDGET ITEM JUSTIFI	CATION (R-2A Exhibit)	DATE February 2000
BUDGET A	CTIVITY Died Res	search	PE NUMBER AND TITLE 0602787A Medical Technology	PROJECT A878
FY 2000	Planned I	Program: (continued)		
•	815	- Complete validation of cold water immersion safety limits students.	s for Ranger training using data from temperatu	re pills collected in free-ranging Ranger
•	1044	 Develop standardized baseline UH-60 simulator, flight pe studies. 	rformance database for application within spati-	al disorientation and sustained operations
•	195	- Small Business Innovative Research/Small Business Tech	nology Transfer Research Programs.	
Total	9267			
FY 2001 I	Planned P	rogram:		
•	2997	- Develop treatment guidance for a field therapy kit for lase	ŭ •	
•	3089	- Validate predictive finite element models of blunt trauma,		
•	1026	- Develop predictive models of head-supported mass and no modeling with in-flight testing.		•
•	2265	- Evaluate the effects of physical fatigue, sleep deprivation, cold exposures.	and other operational stressors on the pathophy	ysiological responses to acute or chronic
•	1265	- Assess the impact of fatigue countermeasures and training	on prevention of spatial disorientation acciden	ts.
Total	10642			
Project A	878	Pao	e 23 of 40 Pages	Exhibit R-2A (PE 0602787A)

	A	ARMY RDT&E BUDGET ITE	EM JUS	TIFICAT	ΓΙΟΝ (R-	-2A Exh	ibit)		DATE Fe	bruary 20	000
BUDGET AC 2 - Appl		search			UMBER AND 02787A		echnolo	gy	PROJECT A879		
		COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate	FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost
A879 Medi	cal Factors	Enhancing Soldier Effectiveness	7759	8019	8438	8725	9015	10214	10739	Continuing	Continuin
		and Justification: This project focuses or									
		sis is on identification of baseline physiolo									
collection of	f rules and	d algorithms for performance degradation i	n multistress	or environm	ents form the	e basis for th	e developme	ent of behav	ioral, trainin	g, pharmacol	logical and
nutritional ("skin-in") interventions to prevent decrements and s	ustain soldie	r performan	ce. Key stres	ssors include	psychologic	cal stress fro	m isolation,	new operation	onal roles,
		nents; inadequate restorative sleep; prolong					reme enviro	nments; des	ynchronizatio	on of biologi	cal
rhythms dur	ring deplo	syments across multiple time zones and night	ht operations	; and therma	al and altitud	e stress.					
FY 1999 A	ccomplis	hmants									
1 1 <i>1))) A</i>		- Demonstrated that low aerobic fitness w	as a sionific	ant risk facto	or for serious	s injury in bo	th men and	women iinor	entry into h	asic combat	training
•	1246	- Developed initial computer models for t									
·	1240	capture system and software analysis prog									
		initial tests to determine optimal time ove						i irequeire y t	ina contrast	ieveis. Com	protou
•	833	- Determined that the thermoregulatory sy						oduction wh	en multiple (rold exposur	es are
•	033	repeated within a day.	stem range	ies as mare	ated by bluif	ung or metal	some mean pr	oduction wil	en manapie v	гога спроваг	CB arc
•	2650	- Discovered problem side effect with Mo	odafinil in av	iators with r	nultiple high	doses.					
•	1736	- Developed and refined integration of SC					ediction of pl	hysiological	responses to	heat and co	ld stress
		application in training and operations for						<i>j.</i>			
Total	7759	5 .									
FY 2000 P	lanned P	rogram:									
•	1346	- Quantify effects of current and develop	mental load-	carrying gear	r, clothing, a	nd individua	l equipment	configured i	for specific s	quad positio	ns on the
		biomechanics and physical performance of	of warfighter	s.					-		
•	1824	- Study the effects of high OPTEMPO/PE	ERSTEMPO	on soldier a	nd unit readi	ness for a wi	de range of	military outo	comes includ	ing marksma	anship,
		soldier retention, and soldier physical and									
•	2616	- Validate models for predicting the water					iting in moui	ntain enviror	nments.		
•	967	- Develop interim Health Hazards Assess			-	•					
•	1139	- Transition caffeine research, including of		ulations, opti	imal dosing,	and effects i	n habituated	and nonhal	bituated user	s, to a caffei	ne product
		and/or guidance for caffeine use in the fie									
	127	- Small Business Innovative Research/Sm	nall Business	Technology	Transfer Re	search Prog	rams.				
•				C.	, iransici ic	35carcii 1105					
• Total	8019			2,	, Transfer ix	oscuren 110g					

	A	RMY RDT&E BUDGET IT	TEM JUSTIFICATION (R-2A Exhibit) PATE February 2000
BUDGET AC 2 - Appl	TIVITY lied Res	search	PE NUMBER AND TITLE 0602787A Medical Tech	nnology A879
FY 2001 P	Planned Pr	ogram:		
•			ators to maximize regional dry heat loss in combination v	with current microclimate cooling techniques.
•	1580		nd body fluid shifts to better predict initial stages of heat	
•	1618	- Demonstrate modafinil efficacy for mi	ilitarily relevant performance sustainment in flight and in	n field environments.
•	1354	- Identify application of objective physicompany-size unit during a real-world r	ological test such as voice stress analysis and pupillome	try to assess military performance including study of
•	2589		ed loads affect the biomechanical and performance para	meters and extend the gait model to accommodate
Total	8438	varying terrain conditions.		
roject A8	270		Page 25 of 40 Pages	Exhibit R-2A (PE 0602787A)

ARMY RDT&E BUDGET I	DATE Fe	February 2000								
BUDGET ACTIVITY 2 - Applied Research		PE NUMBER AND TITLE 0602787A Medical Technology					PROJECT A921			
COST (In Thousands)	FY 1999 Actual	FY 200 Estimat	-	FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost
A921 Ovarian Cancer Research		11	771	0	0	0	0	0	0	11771

Mission Description and Justification: By Congressional direction, the purpose of this appropriation is only for ovarian cancer research.

FY 1999 Accomplishments: Funded within the Defense Health Program in FY 1999.

FY 2000 Planned Program:

- 11454 Determine FY 2000 vision in a meeting in February 2000. Receive and evaluate proposals.
- 317 Small Business Innovative Research/Small Business Technology Transfer Research Programs.

Total 11771

FY 2001 Planned Program: Project not funded in FY 2001. Award proposals from the FY 2000 program.

Project A921 Page 26 of 40 Pages Exhibit R-2A (PE 0602787A)

ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2A Exhibit) PE NUMBER AND TITLE PROJECT PROJECT										
BUDGET ACTIVITY 2 - Applied Research			UMBER AND		echnolo	ду		ROJECT \948		
COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate	FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cos	
A948 Portable Cardiopulmonary Bypass Pump and Oxygenator	1925	0	0	0	0	0	0	0	192	
- Developed a re-usable but ultimately di Total 1925 FY 2000 Planned Program: Project not funded in FY 2000. A FY 2001 Planned Program: Project not funded in FY 2001.	•	•	•	•	ry bypass fu	nd system.				

Item 28

Exhibit R-2A (PE 0602787A)

Project A948

ARMY RDT&E BUDGE	T ITEM JUS	TIFICAT	ION (R-	2A Exh	ibit)		DATE Fe	bruary 20	00
BUDGET ACTIVITY 2 - Applied Research			UMBER AND 02787A		PROJEC* A949				
COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate	FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cos
N949 Advanced Cancer Detection	3374	0	0 0	0	0	C	0	0	337
- Completed scientific peer review Florida on a scientifically meritor Total 3374 Y 2000 Planned Program: Project funded under program: Project not funded in FY	rious proposal for the	balance of	the appropria	ation.	posai in tuii	and workin	g with the Oi	niversity of S	outn

Item 28

Exhibit R-2A (PE 0602787A)

Project A949

ARMY RDT&E BUDGET IT	EM JUS	TIFIC	CAT	TON (R	-2A Exh	ibit)		DATE Fe	bruary 20	000
BUDGET ACTIVITY 2 - Applied Research				PE NUMBER AND TITLE 0602787A Medical Technology					PROJECT A950	
COST (In Thousands)	FY 1999 FY 20 Actual Estim			FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost
A950 Teleradiology	2890		0	0	0	0	0	0	0	289
imaging to be deployed in remote and far-forward locations. A medical studies for interpretation. FY 1999 Accomplishments:	•					•				
Project A950		Page	29 of	f 40 Pages			Exhib	it R-2A (PE	0602787A)	

ARMY RDT&E BUDGET	ITEM JUS	STIFICA	TION (R	-2 Exhi	bit)		DATE Fe	bruary 20	000
BUDGET ACTIVITY 2 - Applied Research		PE N 06 0	PROJECT A951						
COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate	FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost
A951 Diagnostic and Surgical Breast Imaging	1926	0	0	0	0	(0	0	1926
Mission Description and Justification: By Congressional de FY 1999 Accomplishments: • 1926 - Received 2-year funds in January 1999 proposal was recommended for fundin Materiel Command, directed re-comperate - Re-advertised the project on August 2 Total 1926 FY 2000 Planned Program: Project not funded in FY 2000 May 2000 for FY 1999 awards. FY 2001 Planned Program: Project not funded in FY 2001 Planned Program Project not funded in FY 2001 Plann	99. Published a g at programma stition. 26, 1999 and red. Conduct scient	program an atic review. ceived 23 pr	nouncement The Comma oposals by C	in March 19 nding Gener October 1999	999. Receive al of the Uni	ed and peer I	reviewed two	proposals. l Research a	nd
Project A951		Page 30 or	f 40 Pages			Exhil	oit R-2 (PE (0602787A)	

ARMY RDT&E BUDGET ITE	RMY RDT&E BUDGET ITEM JUSTIFICATION (R-2A Exhibit)							February 2000			
BUDGET ACTIVITY 2 - Applied Research	PE NUMBER AND TITLE 0602787A Medical Technology							PROJEC A952			
COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate	FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost		
A952 Musculoskeletal Injuries	1926	5885	0	0	0	0	0	0	7811		

Mission Description and Justification: By Congressional direction, the purpose of this project is to develop initial research models for musculoskeletal injuries.

FY 1999 Accomplishments:

1926 - Evaluated competitive contracts/grants to initiate research on musculoskeletal injuries. Awarded research grants.

Total 1926

FY 2000 Planned Program:

• 5727 - A solicitation for research proposals will be developed and competed.

• 158 - Small Business Innovative Research/Small Business Technology Transfer Research Programs.

Total 5885

FY 2001 Planned Program: Project not funded in FY 2001.

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	P	ARMY RDT&E BUDGET	ITEM JUS	TIFICA	ΓΙΟΝ (R-	-2A Exh	ibit)		DATE Fe	bruary 20	000
BUDGET AC 2 - App		search			NUMBER AND 02787A		echnolo	gy			ROJECT \953
		COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate	FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost
A953 Disa	aster Relief a	and Emergency Medical Services	9630	9809	0	0	0	0	0	0	1943
FY 1999 A Total FY 2000 F Total FY 2001 F	9630 9630 Planned Pr 9545 264 9809	- Continued development of disaster A&M.	elief and emergen h/Small Business	acy and biol	ogical medic	al response c	capability at		·		
Project A9	953			Page 32 o	of 40 Pages			Exhibi	t R-2A (PE	0602787A)	

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ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2A Exhibit)								February 2000		
BUDGET ACTIVITY 2 - Applied Research	PE NUMBER AND TITLE 0602787A Medical Technology					PI A				
COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate		FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost	
A962 Polynitroxylated Hemoglobin	0	19	62 0	0	0	0	0	0	1962	

<u>Mission Description and Justification</u>: By Congressional direction, the purpose of this program is to establish a process to select medical research projects of clear scientific merit and direct relevance to military health including polynitroxylated hemoglobin.

FY 1999 Accomplishments: Project not funded in FY 1999.

FY 2000 Planned Program:

- 1909 Awaiting proposal submission for evaluation to be followed by contract award.
- 53 Small Business Innovative Research/Small Business Technology Transfer Research Programs.

Total 1962

FY 2001 Planned Program: Project not funded in FY 2000.

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-	ARMY RDT&E BUDGET IT	EM JUS	TIFICAT	ΓΙΟΝ (R-	2A Exh	ibit)		DATE Fe	bruary 20	000
BUDGET ACTIVITY 2 - Applied Re	search			UMBER AND 02787A		echnolo	gy	•	ROJECT \963	
	COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate	FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cos
A963 National Medic	al Testbed	0	14714	0	0	0	0	0	0	147
FY 2000 Planned P 14318 396 Total 14714	rogram: - Award cooperative (research) agreement advanced medical (and supporting nonment trauma, and emergency health care in a (- Small Business Innovative Research/Smargram: Project not funded in FY 2000.	edical) techno military/civili	logies, and an) commu	clinical and nity setting.	epidemiolog	ical studies t				

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Project A963

ARMY RDT&E BUDGET	ITEM JUS	ΓΙ <mark>ΓΙ</mark> ΓΑΊ	ΓΙΟΝ (R-	-2A Exh	ibit)		DATE Fe	bruary 20	000
BUDGET ACTIVITY 2 - Applied Research			PE NUMBER AND TITLE 0602787A Medical Technology						ROJECT 4964
COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate	FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cos
A964 Infomatics-based Medical Emergency Tools	0	4414	0	0	0	0	0	0	44
FY 2000 Planned Program: • 4295 - Award cooperative research agreemed architecture that will enhance diagnos • 119 - Small Business Innovative Research Total 4414 FY 2001 Planned Program: Project not funded in FY 2006	ent to conduct ex is, treatment, and /Small Business	d patient ma	ınagement (e	.g., traumatio	c and chemic				and

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Exhibit R-2A (PE 0602787A)

Project A964

A	ARMY RDT&E BUDGE	T ITEM JUS	TIFICAT	ION (R	-2A Exh	ibit)		DATE Fe	bruary 20	000	
BUDGET ACTIVITY 2 - Applied Res	search			UMBER AND 02787A	TITLE Medical T	echnolo	gy		PROJECT A965		
	COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate	FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost	
A965 Eye Research		0	1962	0	0	0	0	0	0	196	
FY 2000 Planned P.	9	ement to conduct exate treatment of, an arch/Small Business	d minimize t	he effects (c	on human bel	navior) of lo		s (e.g., instru	mentation re	search)	

ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2A Exhibit)								February 2000			
BUDGET ACTIVITY 2 - Applied Research	PE NUMBER AND TITLE 0602787A Medical Technology						PROJECT A966				
COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate		FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost		
A966 Blood Research	0	53	395	0 (0	0	0	0	5395		

<u>Mission Description and Justification</u>: By Congressional direction, the purpose of this program is for research into improved blood products and safety in systems compatible with military field use.

FY 1999 Accomplishments: Project not funded in FY 1999.

FY 2000 Planned Program:

- 5250 Awaiting proposal submission for evaluation to be followed by contract award.
- 145 Small Business Innovative Research/Small Business Technology Transfer Research Programs.

Total 5395

FY 2001 Planned Program: Project not funded in FY 2000.

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ARMY RDT&E BUDGET ITEM JUSTIFICATION (R-2A Exhibit)								February 2000			
BUDGET ACTIVITY 2 - Applied Research	PE NUMBER AND TITLE 0602787A Medical Technology								PROJECT A967		
COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate	FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost		
A967 Dye Targeted Laser Fusion	0	294	13 0	0	0	0	0	0	2943		

Mission Description and Justification: By Congressional direction, the purpose of this program is for research into tissue repair systems.

FY 1999 Accomplishments: Project not funded in FY 1999.

FY 2000 Planned Program:

• 2864 - Awaiting proposal submission for evaluation to be followed by contract award.

• 79 - Small Business Innovative Research/Small Business Technology Transfer Research Programs.

Total 2943

FY 2001 Planned Program: Project not funded in FY 2000.

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ARMY RDT&E BUDGE	T ITEM JUS	TIFICAT	ION (R-	2A Exh	ibit)		DATE Fe	bruary 20	000
BUDGET ACTIVITY 2 - Applied Research			UMBER AND TO 1981		echnolo	gy			ROJECT \968
COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate	FY 2001 Estimate	FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cos
A968 Synchrotron-based High Energy Radiation Beam	0	4905	0	0	0	0	0	0	49
application of proton beam radiation therapy (supported b FY 1999 Accomplishments: Project not funded in FY 1 FY 2000 Planned Program:	ement to Loma Lind d switchyard that w awaiting release by arch/Small Business	da Universit ill enable pr OSD for FY	y Medical Ce ecise, extend Y 2000.	enter to cond	luct explorat	ory develop	ment, integra	ation, and	

ARMY RDT&E BUDGET IT	EM JUS	TIFICA	ATION (F	R-2A Exh	ibit)		February 2000			
BUDGET ACTIVITY 2 - Applied Research		E NUMBER AN 1602787A		PROJECT A977						
COST (In Thousands)	FY 1999 Actual	FY 2000 Estimate		FY 2002 Estimate	FY 2003 Estimate	FY 2004 Estimate	FY 2005 Estimate	Cost to Complete	Total Cost	
A977 Emerging Infectious Diseases	0		0 69:	57 0	0	0	0	0	695	

FY 1999 Planned Program: Project not funded in FY 1999.

necessary research to counter the military operational impact of emerging infectious diseases.

FY 2000 Planned Program: Project not funded in FY 2000.

FY 2001 Planned Program:

6957 - Complete applied research to characterize possible tools and components of protective measures against emerging infectious diseases.

Total 6957

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